



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:

Agoston et al.

Serial No. 09/939,208

Filed: August 24, 2001

For: ANTIANGIOGENIC AGENTS

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APPENDIX A

Assistant Commissioner for Patents  
Washington, D.C. 20231  
Sir:

Applicants respectfully request that the Examiner review the below-listed EntreMed patent applications. Please review and initial each listed application and return a copy of the initialed Appendix A to the Applicants.

The serial numbers of the below-listed U.S. patent applications are listed on this form as provided in MPEP §609(D), rather than a PTO-1449 form, to avoid the serial numbers of the pending U.S. patent applications being published on any patent that may issue from this application. Applicants do not waive the confidential nature of any of the listed patent applications.

- \_\_\_\_\_ 1. U.S. Application Serial No. 09/641,327 filed August 18, 2000; Atty. File No. 05213-0730/219693
- \_\_\_\_\_ 2. U.S. Application Serial No. 09/779,331 filed February 8, 2001; Atty. File No. 05213-0731/253406
- \_\_\_\_\_ 3. U.S. Application Serial No. 09/780,650 filed February 12, 2001; Atty. File No. 05213-0493/253692
- \_\_\_\_\_ 4. U.S. Application Serial No. 09/866,279 filed May 25, 2001; Atty. File No. 05213-0412/259097

- \_\_\_\_\_ 5. U.S. Application Serial No. 09/899,702 filed July 5, 2001; Atty. File No. 05213-0910/260981
- \_\_\_\_\_ 6. PCT Application Serial No. PCT/US01/26490 filed August 24, 2001; Atty. File No. 05213-0852WP/263551
- \_\_\_\_\_ 7. U.S. Application Serial No. Not Yet Assigned filed February 15, 2002 by Robert D'Amato entitled "Antiangiogenic Agents and Methods for Treatment of Neovascularization and Related Conditions"; Atty. Docket No. 05213-3000/269288.

Respectfully Submitted,

By: Suzanne Seavello Shope  
Suzanne Seavello Shope  
Reg. No. 37,933

KILPATRICK STOCKTON LLP  
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Our Docket: 05213-0852 (43170-263550)

Substitute for Form 1449/A/PTO <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)				Complete if Known		
				Application No.	09/939,208	
				Filing Date	08/24/2001	
				First Named Inventor	Agoston et al.	
				Group Art Unit	1623 1616	
Examiner Name	Unassigned					
Attorney Docket Number	05213-0852 43170-263550					
Sheet	1	of	16			
<b>U.S. PATENT DOCUMENTS</b>						
Examiner Initials	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number <sup>2</sup>	Kind Code <sup>3</sup> (if known)			
	1	2,584,271		Huffman	02/05/1952	
	2	2,846,453		Hoehn	08/05/1958	
	3	3,410,879		Smith et al.	11/12/1968	
	4	3,470,218		Farah	09/30/1969	
	5	3,492,321		Crabbe	01/27/1970	
	6	3,496,272		Kruger (Identifier only)	02/17/1970	
	7	3,956,348		Hilscher	05/11/1976	
	8	4,172,132		Draper et al.	10/23/1979	
	9	4,212,864		Tax	07/15/1980	
	10	4,307,086		Tax	12/22/1981	
	11	4,522,758		Ward et al.	06/11/1985	
	12	4,743,597		Javitt et al.	05/10/1988	
	13	4,994,443	B1	Folkman et al.	02/19/1991	
	14	5,001,116		Folkman et al.	03/19/1991	
	15	5,135,919		Folkman et al.	08/04/1992	
	16	5,504,074		D'Amato et al.	04/02/1996	
	17	5,521,168		Clark	05/28/1996	
	18	5,621,124		Seitz et al.	04/15/1997	
	19	5,643,900		Fotsis et al.	07/01/1997	
	20	5,661,143		D'Amato et al.	08/26/1997	
	21	5,892,069		D'Amato et al.	04/06/1999	
	22	6,011,023	B1	Clark et al.	01/04/2000	
	23	6,136,992		Ram et al.	10/24/2000	
	24	6,200,966	B1	Stewart	03/13/2001	
Examiner Signature	<i>S. O. [Signature]</i>				Date Considered	2/23/06

<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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				Application No.	09/939,208	
				Filing Date	08/24/2001	
				First Named Inventor	Agoston et al.	
				Group Art Unit	1823/616	
				Examiner Name	Unassigned <i>GAZIS</i>	
Sheet	2	of	16	Attorney Docket Number		05213-0852 43170-263550
<b>U.S. PATENT DOCUMENTS</b>						
Examiner Initials	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number <sup>2</sup>	Kind Code <sup>3</sup> (if known)			
<i>GB</i>	25	6,239,123	B1	Green et al.	05/29/2001	

Examiner Signature	<i>SCD</i>	Date Considered	<i>2/23/06</i>
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				First Named Inventor	Agoston et al.
				Group Art. Unit	1623 1616
				Examiner Name	Unassigned QA21, S
Sheet 3 of 16				Attorney Docket Number 05213-0852 43170-263550	

## FOREIGN PATENT DOCUMENTS

Examiner Initials	Cite No. <sup>1</sup>	Foreign Patent Document			Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T <sup>2</sup>
		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
	26	DE	1907330		Sandoz Ltd. (Abstract only)	10/23/1969		
	27	DE	3625315		Schering A.-G (Abstract only)	01/28/1998		
SC	28	WO	90/15816	A1	The Upjohn Company	12/27/1990		
	29	WO	93/03729		Research Corporation Technologies, Inc.	03/04/1993		
	30	WO	95/04535		The Children's Medical Center Corporation	02/16/1995		
	31	WO	98/40398		Pharm-Eco Laboratories, Inc.	09/17/1998		

Examiner Signature	<i>SC</i>	Date Considered	2/23/06
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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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		First Named Inventor	Agoston et al.
		Group Art Unit	1623/1616
		Examiner Name	Unassigned (GA-2)
Attorney Docket Number	05213-0852 43170-26358		
<b>OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS</b>			
Examiner Initials	Cite No.	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>a</sup>
	32	Author: Title: Lioipristone/(1-[4-(Dimethylamino)phenyl]-17-hydroxy-17-(3-hydroxy-1-propenyl)estra-4,9-diene-3-one; AK 98734 Publ: Dict. of Drugs (1990), Dict. of Steroids (1991), Dict. of Org. Vol/Iss: Pages: Date: Cmpds (6th Ed) (1996), Dict. of Pharm. Agents (1997)	
	33	Author: Title: (paragraphs 583-584) Publ: The Merck Index 11th Edition Vol/Iss: Pages: 88 Date: 1989	
	34	Author: ADAMS, E.F. et al. Title: Steroidal regulation of oestradiol-17B dehydrogenase activity of the human breast cancer cell line MCF-7 Publ: Journal of Endocrinology Vol/Iss: 188 (1) Pages: 149-154 Date: 07/1988	
	35	Author: AIZU-YOKOTA et al. Title: Natural Estrogens Induce Modulation of Microtubules in Chinese Hamster V79 Cells in Culture Publ: Cancer Research Vol/Iss: 55 Pages: 1863-1868 Date: 05/01/95	
	36	Author: ATTALIA et al. Title: 2-Methoxyestradiol Arrests Cells in Mitosis without Depolarizing Tubulin Publ: Biochemical and Biophysical Research Communications Vol/Iss: 228 Pages: 467-473 Date: 1996	
	37	Author: AYALA et al. Title: The Induction of Accelerated Thymic Programmed Cell Death During Polymicrobial Sepsis: Control by Corticosteroids but not Tumor Necrosis Factor (Abstract only) Publ: Shock Vol/Iss: 3 (4) Pages: 259-267 Date: 04/1995	
	38	Author: BANIK et al. Title: Orally Active Long-Acting Estrogen (3-(2-propynyloxy)-estra-1,3,5,(10)-triene-17. beta.-ol trimethylacetate) (Identifier only) Publ: Steroids Vol/Iss: 16 (3) Pages: 289-296 Date: 1970	

Examiner Signature	<i>S. Q. 3</i>	Date Considered	2/23/06
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<sup>a</sup>Unique citation designation number. <sup>b</sup>See attached Kinds of U.S. Patent Documents. <sup>c</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>d</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>e</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>f</sup>Applicant is to place a check mark here if English language translation is attached.

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<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)		Application No.	09/939,208
		Filing Date	08/24/2001
		First Named Inventor	Agoston et al.
		Group Art Unit	1623 1616
		Examiner Name	Unassigned 8/21/05
Sheet 5 of 16	Attorney Docket Number	05213-0852 43170-263550	
<b>OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS</b>			
Examiner Initials	Cite No.	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T*
	39	Author: BARDON et al. Title: Steroid Receptor-Mediated Cytotoxicity of an Antiestrogen and an Antiprogesterin in Breast Cancer Cells (Abstract only) Publ: Cancer Research Vol/Iss: 47 (5) Pages: 1441-1448 Date: 03/01/87	
	40	Author: BHAT et al. Title: Estradiol-Induced Mitotic Inhibition in the Bursa of Fabricius of Male Domestic Duckling Publ: Mikroskopie Vol/Iss: 39 Pages: 113-117 Date: 05/1982	
	41	Author: BLAGOSKLONNY et al. Title: Raf-1/bcl-2 Phosphorylation: A Step from Microtubule Damage to Cell Death Publ: Cancer Research Vol/Iss: 57 Pages: 130-135 Date: 01/01/97	
	42	Author: BLICKENSTAFF et al. Title: Estrogen-Catharanthus (Vinca) Alkaloid Conjugates Publ: Cytotoxic Estrogens in Hormone Receptive Tumors Vol/Iss: Pages: 89-105 Date: 1980	
	43	Author: BOYE et al. Title: 185. Deaminocolchinyll Methyl Ether. Synthesis from 2,3,4,4'-Tetramethoxybiphenyl -2-carbaldehyde. Comparison of Antitubulin Effects of Publ: Helvetica Chimica Acta Vol/Iss: 72 Pages: 1690-1696 Date: 1989	
	44	Author: BRODIE, A.M. Title: Aromatase Inhibitors in the Treatment of Breast Cancer (Abstract only) Publ: Journal of Steroid Biochemistry and Molecular Biology Vol/Iss: 49 (4-6) Pages: 281-287 Date: 06/1994	
	45	Author: BROSENS et al. Title: Comparative Study of the Estrogenic Effect of Ethinylestradiol and Mestranol on the Endometrium Publ: Laboratory for Gynecological Physiopathology Vol/Iss: 14 (6) Pages: 679-685 Date: 12/01/76	

Examiner Signature	<i>S. B. 23</i>	Date Considered	2/23/06
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\*Unique citation designation number. \*See attached Kinds of U.S. Patent Documents. \*Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). \*For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. \*Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. \*Applicant is to place a check mark here if English language translation is attached.

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<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)		Application No.	09/939,208
		Filing Date	08/24/2001
		First Named Inventor	Agoston et al.
		Group Art Unit	3623 1616
		Examiner Name	Unassigned BPAZ, 8/21/01
Sheet 6 of 16		Attorney Docket Number	05213-0852 43170-263550
<b>OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS</b>			
Examiner Initials	Date No.	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	
	46	Author: CASTAGNETTA, L. et al. Title: Simple Approach to Measure Metabolic Pathways of Steroids in Living Cells Publi: Journal of Chromatography Vol/Iss: 572 Pages: 25-39 Date: 12/06/91	
	47	Author: CHASSEROT-GOLAZ et al. Title: Biotransformation of 17.beta.-hydroxy-11.beta.-(4-dimethylaminophenyl) 17.alpha.1-propynyl-estra-4,9-diene-3-one (RU486) in Rat Hepatoma Variants (Identifier only) Publi: Biochemical Pharmacology Vol/Iss: 46 (11) Pages: 2100-2103 Date: 01/01/93	
	48	Author: CHEN et al. Title: A New Synthetic Route to 2- and 4-Methoxyestradiols by Nucleophilic Substitution Publi: Steroids Vol/Iss: 47 (1) Pages: 63-66 Date: 01/1986	
SC	49	Author: CHEN et al. Title: Synthesis of 11.beta.-(4-dimethylaminophenyl)-17.beta.-hydroxy-17.alpha.-(1-propyl) estr-4, 9-dien-3-one (RU486) (Identifier only) Publi: Nanjing Yaoxueyuan Xuebao Vol/Iss: 17 (4) Pages: 282-285 Date: 1986	
	50	Author: COHEN et al. Title: Novel Total Synthesis of (+)-Estrone 3-Methyl Ether, (+)-13B-Ethyl-3-methoxygona-1,3,5(10-triene-17-one, and (+)-Equilenin 3-Methyl Ether Publi: The Journal of Organic Chemistry Vol/Iss: 6 Pages: 681-685 Date: 03/21/75	
	51	Author: COLLINS et al. Title: The Structure and Function of Estrogens. XI* Synthesis of (+/-)-7(8-11a) abeo-Estradiol and its 9,11-Didehydro Derivative Publi: Aust. Journal of Chemistry Vol/Iss: 45 Pages: 71-97 Date: 1992	
	52	Author: CRUM, R. et al. Title: A New Class of Steroids Inhibits Angiogenesis in the Presence of Heparin or a Heparin Fragment Publi: SCIENCE Vol/Iss: 230 Pages: 1375-1378 Date: 12/20/85	

Examiner Signature	SC	Date Considered	2/23/06
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STATEMENT BY APPLICANT

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Sheet 7 of 16

Application No.	09/939,208
Filing Date	08/24/2001
First Named Inventor	Agoston et al.
Group Art Unit	1623 1616
Examiner Name	Unassigned
Attorney Docket Number	05213-0852 43170-26359

## OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS

Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
	53	Author: CUMMINGS et al. Title: Apoptosis Publ: The American Journal of Surgical Pathology Vol/Iss: 21 (1) Pages: 88-101 Date: 1997	
	54	Author: CUSHMAN et al. Title: Synthesis, Antitubulin and Antimitotic Activity, and Cytotoxicity of Analogs of 2-Methoxyestradiol, an Endogenous Mammalian Metabolite of Estradiol that Publ: Journal of Medical Chemistry Vol/Iss: 38 (12) Pages: 2041-2049 Date: 1995	
	55	Author: CUSHMAN et al. Title: Synthesis of Analogs of 2-Methoxyestradiol with Enhanced Inhibitory Effects of Tubulin Polymerization and Cancer Cell Growth Publ: Journal of Medical Chemistry Vol/Iss: 40 (15) Pages: 2323-2334 Date: 1997	
	56	Author: D'AMATO et al. Title: 2-Methoxyestradiol, an Endogenous Mammalian Metabolite, Inhibits Tubulin Polymerization by Interacting at the Colchicine Site Publ: Proceedings of the National Academy of Science USA Vol/Iss: 91 Pages: 3964-3968 Date: 04/1994	
	57	Author: DVIR et al. Title: Thin-layer Chromatography of DANSYL-oestrogens Publ: Journal of Chromatography Vol/Iss: 52 Pages: 505-506 Date: 11/04/70	
	58	Author: EPE et al. Title: Microtubular Proteins as Cellular Targets for Carcinogenic Estrogens and Other Carcinogens Publ: Mechanisms of Chromosome Distribution and Aneuploidy Vol/Iss: Pages: 345-351 Date: 1989	
	59	Author: EVANS et al. Title: A Convergent Total Synthesis of (+)- Colchicine and (+)-Desacetamidolsocolchicine Publ: Journal of the American Chemical Society Vol/Iss: 103 Pages: 5813-5821 Date: 09/23/81	

Examiner  
Signature

S. G. 20

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Sheet

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16

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Examiner Name	Unassigned QA21, S.
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	60	Author: FISHMAN, J. Title: Synthesis of 2-Methoxyestrogens Publ: Journal of the American Chemical Society Vol/Iss: 80 Pages: 1213-1216 Date: 03/05/98	
	61	Author: FITZGERALD Title: Molecular Features of Colchicine Associated with Antimitotic Activity and Inhibition of Tubulin Polymerization Publ: Biochemical Pharmacology Vol/Iss: 25 Pages: 1383-1387 Date: 06/15/76	
	62	Author: FOTSIS et al. Title: The Endogenous Oestrogen Metabolite 2-Methoxyoestradiol Inhibits Angiogenesis and Suppresses Tumour Growth Publ: Nature Vol/Iss: 368 Pages: 237-239 Date: 03/17/94	
	63	Author: GETAHUN et al. Title: Synthesis of Alkoxy-Substituted Diaryl Compounds and Correlation of Ring Separation with Inhibition of Tubulin Polymerization: Differential Enhancement of Inhibitory Effects Under Suboptimal Polymerization Publ: Journal of Medical Chemistry Vol/Iss: 35 (6) Pages: 1058-1067 Date: 03/20/92	
	64	Author: GROSS et al. Title: Inhibition of Tumor Growth, Vascularization, and Collagenolysis in the Rabbit Cornea by Medroxyprogesterone Publ: Proceedings of the National Academy of Science USA Vol/Iss: 78 (2) Pages: 1176-1180 Date: 02/1981	
	65	Author: GUANGRONG et al. Title: Effect of Components of Growth Ether Copper(I)iodide Mixed Catalyst on Nucleophilic Substitution of Bromoestrogen (Abstract No. 195225) Publ: Chemical Abstracts Vol/Iss: 111 (21) Pages: 818, Column 1 Date: 11/20/89	
	66	Author: HALDAR et al. Title: Bc12 is the Guardian of Microtubule Integrity Publ: Cancer Research Vol/Iss: 57 Pages: 229-233 Date: 01/15/97	

Examiner  
Signature

S. Q. [Signature]

Date  
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Sheet

10

of

16

## Complete if Known

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09/939,208

Filing Date

08/24/2001

First Named Inventor

Agoston et al.

Group Art Unit

1623 1616

Examiner Name

Unassigned

Attorney Docket Number

05213-0852 43170-263550

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	74	Author: KLAUBER et al. Title: Inhibition of Angiogenesis and Breast Cancer in Mice by the Microtubule Inhibitors 2-Methoxyestradiol and Taxol Publ: Cancer Research Vol/Iss: 57 Pages: 81-88 Date: 01/01/97	
	75	Author: LEBRAS, J. et al. Title: Activation and Regioselective Ortho-Functionalization of the A-Ring of B-Estradiol Promoted by "Cp*Ir": An Efficient Organometallic Procedure for the Synthesis of 2-Methoxyestradiol Publ: Organometallics Vol/Iss: 16 Pages: 1765-1771 Date: 1997	
SB	76	Author: LIN et al. Title: Interactions of Tubulin with Potent Natural and Synthetic Analogs of the Antimitotic Agent Combretastatin: A Structure-Activity Study Publ: Molecular Pharmacology Vol/Iss: 34 (2) Pages: 200-208 Date: 08/1988	
	77	Author: LINCOLN et al. Title: Conformation of Thiocolchicine and Two B-Ring-Modified Analogues Bound to Tubulin Studied with Optical Spectroscopy Publ: Biochemistry Vol/Iss: 30 (5) Pages: 1179-1187 Date: 02/05/91	
	78	Author: LOTTERING et al. Title: Effects of the 17 $\beta$ -Estradiol Metabolites on Cell Cycle Events in MCF-7 Cells Publ: Cancer Research Vol/Iss: 52 Pages: 5926-5932 Date: 11/01/92	
	79	Author: LOTTERING et al. Title: 17 $\beta$ -Estradiol Metabolites Affect Some Regulators of the MCF-7 Cell Cycle Publ: Cancer Letters Vol/Iss: 110 Pages: 181-186 Date: 1996	
	80	Author: MAYOL et al. Title: Ethynylestradiol-Induced Cell Proliferation in Rat Liver Involvement of Specific Populations of Hepatocytes (Abstract only) Publ: Carcinogenesis Vol/Iss: 12 (12) Pages: 2381-2388 Date: 1992	

Examiner Signature

S. Q. 2

Date

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2/23/06

<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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# INFORMATION DISCLOSURE STATEMENT BY APPLICANT

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Sheet

9

of

16

Application No.

09/939,208

Filing Date

08/24/2001

First Named Inventor

Agoston et al.

Group Art Unit

1625/1616

Examiner Name

Unassigned BAZIS

Attorney Docket Number

05213-0852 43170-263550

## OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS

Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
	67	Author: HAMEL et al. Title: Interactions of 2-Methoxyestradiol, an Endogenous Mammalian Metabolite, with Unpolymerized Tubulin and with Tubulin Polymers Publ: Biochemistry Vol/Iss: 35 (4) Pages: 1304-1310 Date: 1996	
	68	Author: HARTLEY-ASP et al. Title: Diethylstilbestrol Induces Metaphase Arrest and Inhibits Microtubule Assembly Publ: Mutation Research Vol/Iss: 143 (4) Pages: 231-235 Date: 08/1985	
503	69	Author: HE et al. Title: A Versatile Synthesis of 2-Methoxyestradiol, an Endogenous Metabolite of Estradiol which Inhibits Tubulin Polymerization by Binding to the Colchicine Binding Site Publ: Bioorganic & Medicinal Chemistry Letters Vol/Iss: 4 (14) Pages: 1724-1728 Date: 1994	
	70	Author: HUBER et al. Title: Tubulin Binding of Conformationally Restricted Bis-Aryl Compounds Publ: Bioorganic & Medicinal Chemistry Letters Vol/Iss: 1 (5) Pages: 243-246 Date: 1991	
	71	Author: JOSEFSSON et al. Title: Suppression of Type II Collagen-Induced Arthritis by the Endogenous Estrogen Metabolite 2-Methoxyestradiol Publ: Arthritis & Rheumatism Vol/Iss: 40 (1) Pages: 154-163 Date: 01/1997	
	72	Author: KABARITY et al. Title: Further Investigations on the cytological effects of some contraceptives Publ: Mutation Research Vol/Iss: 135 Pages: 181-188 Date: 1984	
	73	Author: KELLY et al. Title: The Stimulation of Prostaglandin Production by Two Antiprogesterone Steroids in Human Endometrial Cells (Abstract only) Publ: Journal of Clinical Endocrinology Metabolism Vol/Iss: 62 (6) Pages: 1116-1123 Date: 06/1986	

Examiner  
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503

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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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16

## Complete if Known

Application No.	09/939,208
Filing Date	08/24/2001
First Named Inventor	Agoston et al.
Group Art Unit	1622 1616
Examiner Name	Unassigned GAZL S
Attorney Docket Number	05213-0852 43170-263550

**OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS**

Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
	81	Author: MEIKRANTZ et al. Title: Apoptosis and the Cell Cycle Publ: Journal of Cellular Biochemistry Vol/Iss: 58 Pages: 160-174 Date: 06/1995	
	82	Author: MILLER et al. Title: Synthesis and Structure-Activity Profiles of A-Homoestrans, the Estratropenes Publ: Journal of Medicinal Chemistry Vol/Iss: 40 Pages: 3836-3841 Date: 1997	
SB	83	Author: MORGAN et al. Title: Calcium and Oestrogen Interactions upon the Rat Thymic Lymphocyte Plasma Membrane Publ: Biochemical and Biophysical Research Communications Vol/Iss: 72 (2) Pages: 663-672 Date: 09/20/76	
	84	Author: MUKHOPADHYAY et al. Title: Induction of Apoptosis in Human Lung Cancer Cells after Wild-Type p53 Activation by Methoxyestradiol Publ: Oncogene Vol/Iss: 14 Pages: 379-384 Date: 1997	
	85	Author: MUKUNDAN et al. Title: Liver Regeneration in Oral Contraceptive Treated Female Rats - Effects of Moderate Malnutrition Publ: Hormone and Metabolic Research Vol/Iss: 16 Pages: 641-645 Date: 12/1984	
	86	Author: NAKAGAWA-YAGI et al. Title: The Endogenous Estrogen Metabolite 2-Methoxyestradiol Induces Apoptotic Neuronal Cell Death <i>In Vitro</i> Publ: Life Sciences Vol/Iss: 58 (17) Pages: 1461-1467 Date: 1996	
	87	Author: NAKAMURA et al. Title: Studies on the Total Synthesis of dl-Colchicine. I. Synthesis of 3-Hydroxy-9, 10, 11-trimethoxy-1,2,3,4,6,7-hexahydro-5H-dibenzo[a,c] cycloheptatrien-5-one Publ: Chemical and Pharmaceutical Bulletin Vol/Iss: 10 Pages: 281-290 Date: 1962	

Examiner Signature

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Date Considered

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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)		Application No.	09/939,208
		Filing Date	08/24/2001
		First Named Inventor	Agoston et al.
		Group Art Unit	1625 1616
		Examiner Name	Unassigned SA215
Sheets	13 of 16	Attorney Docket Number	05213-0852 43170-263550
<b>OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS</b>			
Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
	95	Author: POLI et al. Title: Tumor Necrosis Factor $\alpha$ Functions in an Autocrine Manner in the Induction of Human Immunodeficiency Virus Expression Publ: Proceedings of the National Academy of Science USA Vol/Iss: 87 Pages: 782-785 Date: 01/1990	
	96	Author: RAVINDRA, R. Title: Effect of Estradiol on the <i>in vitro</i> Assembly of Rat Brain Tubulin Publ: Journal of Indian Institute of Science Vol/Iss: 64 (3) Pages: 27-35 Date: 03/1983	
503	97	Author: SAKAKIBARA et al. Title: Effects of Diethylstilbestrol and its Methyl Ethers on Aneuploidy Induction and Microtubule Distribution in Chinese Hamster V79 cells Publ: Mutation Research Vol/Iss: 263 Pages: 269-276 Date: 08/1991	
	98	Author: SATO et al. Title: Effect of Estradiol and Ethynylestradiol on Microtubule Distribution in Chinese Hamster V79 Cells Publ: Chemical and Pharmaceutical Bulletin Vol/Iss: 40 (1) Pages: 182-184 Date: 01/1992	
	99	Author: SATO et al. Title: Disruptive Effect of Diethylstilbestrol on Microtubules Publ: Gann Vol/Iss: 75 Pages: 1046-1048 Date: 12/1984	
	100	Author: SAWADA et al. Title: Colchicine-Like Effect of Diethylstilbestrol (DES) on Mammalian Cells in Vitro Publ: Mutation Research Vol/Iss: 57 Pages: 175-182 Date: 05/1978	
	101	Author: SEEGERs et al. Title: Cyclic-AMP and Cyclic-GMP Production in MCF-7 Cells Exposed to Estradiol-17 Beta, Catecholestrogens and Methoxy-Estrogens in MCF-7 Cells (Meeting Abstract only) Publ: Joint MCI-1st Symposium, Third 1st International Symposium, Biology and Therapy of Breast Cancer Vol/Iss: Pages: Date: 09/25/89	

Examiner Signature	<i>S. O'Brien</i>	Date Considered	2/23/06
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Sheets

12

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16

## Complete if Known

Application No.	09/939,208
Filing Date	08/24/2001
First Named Inventor	Agoston et al.
Group Art Unit	1625 1616
Examiner Name	Unassigned 67A21,5
Attorney Docket Number	05213-0852 43170-263550

**OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS**

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	88	Author: NAMBARA et al. Title: Studies on Steroid Conjugates. III. New Synthesis of 2-Methoxyestrogens Publ: Chem. Pharm. Bulletin Vol/Iss: 18 (3) Pages: 474-480 Date: 03/1970	
	89	Author: NAPOLITANO et al. Title: 11 Beta-Substituted Estradiol Derivatives. 2. Potential Carbon-11 and Iodine-Labeled Probes for the Estrogen Receptor (Abstract only) Publ: Journal of Medical Chemistry Vol/Iss: 38 (14) Pages: 2774-2779 Date: 07/07/95	
	90	Author: NISHIGAKI et al. Title: Anti-Proliferative Effect of 2-Methoxyestradiol on Cultured Smooth Muscle Cells from Rabbit Aorta Publ: Atherosclerosis Vol/Iss: 113 Pages: 167-170 Date: 1995	
	91	Author: OCHS et al. Title: Effect of Tumor Promoting Contraceptive Steroids on Growth and Drug Metabolizing Enzymes in Rat Liver (Abstract only) Publ: Cancer Research Vol/Iss: 46 (3) Pages: 1224-1232 Date: 1986	
	92	Author: OPPOLZER et al. Title: 177. The Enantioselective Synthesis of (+)-Estradiol from 1,3-Dihydrobenzo[c] thiophene-2,2-dioxide by Successive Thermal SO <sub>2</sub> -Extrusion and Cycloaddition Reactions Publ: Helvetica Chimica Acta Vol/Iss: 63 Pages: 1703-1707 Date: 1980	
	93	Author: PARTHASARATHY et al. Title: Antioxidant: A New Role for RU-486 and Related Compounds (Abstract only) Publ: Journal of Clinical Investigation Vol/Iss: 94 (5) Pages: 1990-1995 Date: 11/1994	
	94	Author: PAULL et al. Title: Identification of Novel Antimitotic Agents Acting at the Tubulin Level by Computer-assisted Evaluation of Differential Cytotoxicity Data Publ: Cancer Research Vol/Iss: 52 Pages: 3892-3900 Date: 07/15/92	

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S. [Signature]

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Sheet

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16

Application No.

09/939,208

Filing Date

08/24/2001

First Named Inventor

Agoston et al.

Group Art Unit

1623 1616

Examiner Name

Unassigned QAZI, S

Attorney Docket Number

05213-0852 43170-263550

## OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS

Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published			T <sup>2</sup>
	102	Author: SEEGER, J.C. et al. Publ: <u>Journal of Steroid Biochemistry</u>	Title: The Cytotoxic Effects of Estradiol-17B, Catecholestradiols and Methoxyestradiols on Dividing MCF-7 and HeLa Cells	Vol/Iss: 32 (6) Pages: 797-809 Date: 1989	
	103	Author: SHARP et al. Publ: <u>Carcinogens</u>	Title: Diethylstilbestrol: the Binding and Effects of Diethylstilboestrol upon the Polymerisation and Depolymerisation of Purified Microtubule Protein in vitro	Vol/Iss: 6 (6) Pages: 865-871 Date: 06/1985	
	104	Author: SPICER et al. Publ: <u>Molecular and Cellular Endocrinology</u>	Title: Catecholestrogens Inhibit Proliferation and DNA Synthesis of Porcine Granulosa Cells in Vitro: Comparison with Estradiol, 5 $\alpha$ -dihydrotestosterone, Gonadotropins and Catecholamines	Vol/Iss: 64 Pages: 119-126 Date: 1989	
	105	Author: STERNLICHT et al. Publ: <u>The Journal of Biological Chemistry</u>	Title: Colchicine Inhibition of Microtubule Assembly via Copolymer Formation	Vol/Iss: 254 (20) Pages: 10540-10550 Date: 10/25/79	
	106	Author: SUN et al. Publ: <u>Journal of Medicinal Chemistry</u>	Title: Antitumor Agents. 139. Synthesis and Biological Evaluation of Thiocolchicine Analogs 5,6-Dihydro-6(S)-(acyloxy)-and 5,6-Dihydro-6(S)-[(acyloxy)methyl]-1,2,3-	Vol/Iss: 36 Pages: 544-551 Date: 03/05/93	
	107	Author: SUNAGAWA et al. Publ: <u>Chem. Pharm. Bulletin</u>	Title: Synthesis of Colchicine; Synthesis of d,l'-Demethoxydeoxy-hexahydrocolchicine	Vol/Iss: 9 Pages: 81-83 Date: 1961	
	108	Author: TERANISHI, M. et al. Publ: <u>Chemical and Pharmaceutical Bulletin</u>	Title: Methylation of Catechol Estrogen with Diazomethane	Vol/Iss: 31 (9) Pages: 3309-3314 Date: 09/1983	

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Sabiha Qaz

Date

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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.



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Sheet

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16

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Application No.

09/939,208

Filing Date

08/24/2001

First Named Inventor

Agoston et al.

Group Art Unit

1623/1616

Examiner Name

Unassigned

Attorney Docket Number

05213-0852 43170-263550

**OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS**

Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
	109	Author: TISHLER et al. Title: Microtubule-Active Drugs Taxol, Vinblastine, and Nocodazole Increase the Levels of Transcriptionally Active p53 Publ: Cancer Research Vol/Iss: 55 Pages: 6021-6025 Date: 12/15/95	
	110	Author: TSUTSUI et al. Title: Comparison of Human Versus Syrian Hamster Cells in Culture for Induction of Mitotic Inhibition, Binucleation and Multinucleation, Following Treatment with Four Aneuploidogens Publ: Toxicology in Vitro Vol/Iss: 4 (1) Pages: 75-84 Date: 1990	
	111	Author: UTNE et al. Title: The Synthesis of 2- and 4-Fluoroestradiol Publ: Journal of Organic Chemistry Vol/Iss: 33 (6) Pages: 2469-2473 Date: 06/1968	
	112	Author: VAN GEERESTEIN et al. Title: Structure of 11.beta.-(4-(dimethylamino)phenyl)-17.beta.-hydroxy-17.alpha.-(2-propenyl)estra-4,9-dien-3-one (Identifier only) Publ: Acta Crystall Ogr., Sect. C: Cryst. Struct. Commun. Vol/Iss: C43 (2) Pages: 319-322 Date: 1987	
	113	Author: VAN TAMELEN et al. Title: The Synthesis of Colchicine Publ: Tetrahedron Vol/Iss: 14 Pages: 8-34 Date: 09/1961	
	114	Author: WANG, Z. et al. Title: An Optimized Synthesis of 2-Methoxyestradiol, a Naturally Occurring Human Metabolite with Anticancer Activity Publ: Synth. Commun. Vol/Iss: 28 (23) Pages: 4431-4437 Date: 1998	
	115	Author: WHEELER et al. Title: Mitotic Inhibition and Aneuploidy Induction by Naturally Occurring and Synthetic Estrogens in Chinese Hamster Cells in Vitro Publ: Mutation Research Vol/Iss: 171 Pages: 31-41 Date: 1986	

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Date

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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)		Application No.	09/939,208
		Filing Date	08/24/2001
		First Named Inventor	Agoston et al.
		Group Art Unit	1623
		Examiner Name	Unassigned
Attorney Docket Number	05213-0852 43170-26880		
Sheet 16 of 16			
<b>OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS</b>			
Examiner Initials	Cite No.	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T*
S. O. Z.	116	Author: WHEELER et al. Title: Mitotic Inhibition and Chromosome Displacement Induced by Estradiol in Chinese Hamsters Cells Publ: Cell Motility and the Cytoskeleton Vol/Iss: 7 (3) Pages: 235-247 Date: 1987	
	117	Author: YUE et al. Title: 2-Methoxyestradiol, an Endogenous Estrogen Metabolite, Induces Apoptosis in Endothelial Cells and Inhibits Angiogenesis: Possible Role for Stress-Activated Protein Kinase Signaling Pathway and Fas Expression Publ: Molecular Pharmacology Vol/Iss: 51 Pages: 951-952 Date: 1997	

Examiner Signature	S. O. Z.	Date Considered	2/23/06
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\*Unique citation designation number. \*See attached Kinds of U.S. Patent Documents. \*Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). \*For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. \*Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. \*Applicant is to place a check mark here if English language translation is attached.

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PATENTS

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:

Agoston et al.

Serial No. 09/939,208

Filed: August 24, 2001

For: Antiangiogenic Agents

Group Art Unit: 1616

Examiner: Sabiha Naim Qazi

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Assistant Commissioner for Patents  
Washington, DC 20231

*No page 5 in IDS*

Sir:

The citation of information on the attached Form PTO/SB/08, "List of Art Cited by Applicant" is made pursuant to 37 C.F.R. §§ 1.97 and 1.98. A copy of each listed item is enclosed. In reference to 37 C.F.R. § 1.97(b), this Supplemental Information Disclosure Statement is being filed before the mailing of a first Office action on the merits. Applicants respectfully request that this information be considered by the Office.

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: Assistant Commissioner for Patents, Washington, DC 20231, on May 8, 2002.

*Suzanne Seavello Shope*  
Suzanne Seavello Shope - Reg. No. 37,933

The citation of this information does not constitute an admission that any of the materials are available as a reference or of priority, or a waiver of any right applicant may have under applicable statutes, Rules of Practice in patent cases, or otherwise.

Respectfully submitted,


*Suzanne Seavello Shope*

By: Suzanne Seavello Shope  
Reg. No. 37,933

KILPATRICK STOCKTON LLP  
1100 Peachtree Street  
Suite 2800  
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Our Docket No: 05213-0852 (43170-263550)

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				Application No.		09/939,208		
				Filing Date		08/24/2001		
				First Named Inventor		Agoston		
				Group Art Unit		1616		
				Examiner Name		Sabiha Naim Qazi		
Sheet	1	of	3	Attorney Docket Number				05213-0852 (43170-263550)
<b>FOREIGN PATENT DOCUMENTS</b>								
Examiner Initials	Cite No. <sup>1</sup>	Foreign Patent Document			Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T <sup>2</sup>
		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
SKZ	1	JP	63090763	A	Research Development Corp. of Japan (Abstract only)	04/21/1988		N/A

Examiner Signature		Date Considered	2/23/06
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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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		Group Art Unit	1616
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Sheet	2	of	3
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	2	Author: ARNOLDI et al. Title: Sweet Isovanyl Derivatives: Synthesis and Structure-Taste Relationships of Conformationally Restricted Analogs (abstract only) Publ: Journal of Agric. Food Chem. Vol/Iss: 46(10) Pages: 4002-4010 Date: 1998	
	3	Author: AUDIER et al. Title: Orientation de la fragmentation en spectrometrie de masse par introduction de groupements fonctionnels. VII. - Etihylenecetals de ceto-2 steroides Publ: Bulletin De La Societe Chimique De France Vol/Iss: 10 Pages: 3088-3090 Date: 1965	
5092	4	Author: CAMBIE et al. Title: Aromatic Steroids. Part II. Chromium Trioxide Oxidation of Some Oestra-1,3-5(10)-trienes Publ: Journal of the Chemical Society Vol/Iss: 9 Pages: 1234-1240 Date: 1969	
	5	Author: FETIZON et al. Title: Synthesis of 2-keto steroids (abstract only) Publ: Bull. Soc. Chim. FR. Vol/Iss: 8 Pages: 3301-3306 Date: 1968	
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	7	Author: LIMANTSEV et al. Title: Effect of some estrogen structural analogs on the development of the mouse embryo (abstract only) Publ: Akush Jinekol. Vol/Iss: 6 Pages: 55-56 Date: 1982	
	8	Author: MILLER, Thomas Title: Tubulin as a Therapeutic Target (Abstract only) Publ: Dissertations Abstracts International Vol/Iss: 5907B Pages: 3454 Date: 1998	

Examiner Signature	<i>S. Qazi</i>	Date Considered	2/23/06
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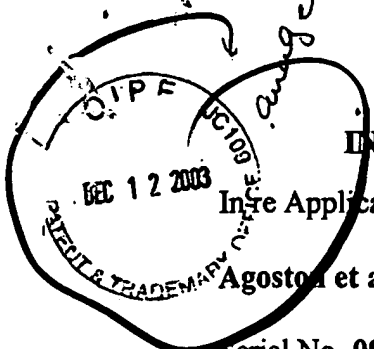
\*Unique citation designation number. \*See attached Kinds of U.S. Patent Documents. \*Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). \*For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. \*Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. \*Applicant is to place a check mark here if English language translation is attached.

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				Application No.	09/939,208
				Filing Date	08/24/2001
				First Named Inventor	Agoston
				Group Art Unit	1616
				Examiner Name	Sabiha Naim Qazi
Sheet	3	of	3	Attorney Docket Number	05213-0852 (43170-263550)
OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS					
Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published			T <sup>2</sup>
	9	Author: NUMAZAWA et al. Title: Efficient Synthesis of 2-Methoxy- and 4-Methoxy-Estrogens Publ: Journal of the Chemical Society <i>Incomplete</i> Vol/Iss: Pages: 533-534 Date: 01/01/83			
<i>SOZ</i>	10	Author: RAO et al. Title: Structural Specificity of Estrogens in the Induction of Mitotic Chromatid Non-Disjunction in Hela Cells Publ: Experimental Cell Research Vol/Iss: 48 Pages: 71-81 Date: 1967			
	11	Author: RAO et al. Title: A Novel, Two-Step Synthesis of 2-Methoxyestradiol Publ: Synthesis <i>Incomplete</i> Vol/Iss: Pages: 168-169 Date: 03/01/77			
<i>SOZ</i>	12	Author: SAKAKIBARA, KYOICHI Title: 2-Hydroxy-1,3,5(10)-estratriene derivatives (abstract only) (Identifier: XP-002186126) Publ: Chemical Abstracts Vol/Iss: 60(1) Pages: Date: 01/06/64			

Examiner Signature	<i>SOZ</i>	Date Considered	2/23/06
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1616  
PATENTS

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:

Agoston et al.

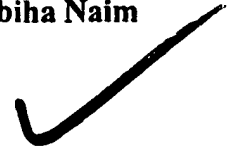
Serial No. 09/939,208

Filed: August 24, 2001

For: Antiangiogenic Agents

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) Art Unit: 1616  
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) Examiner: Qazi, Sabiha Naim  
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SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

Sir:

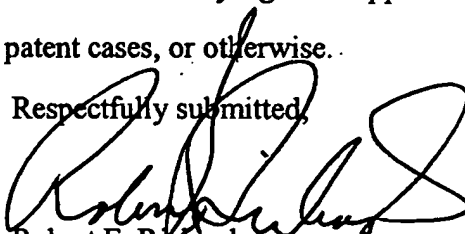
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Respectfully submitted,

  
Robert E. Richards  
Reg. No. 29,105

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Our Docket: 05213-0852 (43170-263550)

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12/16/2003 ZJUHR1 00000027 09939208

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				Application No.	09/939,208
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				First Named Inventor	Gregory E. Agoston et al.
				Group Art Unit	1616
				Examiner Name	Qazi, S.
Sheet	1	of	15	Attorney Docket Number	05213-0852 (43170-263550)

## U.S. PATENT DOCUMENTS

Examiner Initials	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number <sup>2</sup>	Kind Code <sup>3</sup> (if known)			
	1	4,552,758		Murphy et al.	11/12/1985	
	2	4,808,402		Leibovich et al.	02/28/1989	
	3	5,629,340	A	Kuwano et al.	05/13/1997	
	4	5,639,725		O'Reilly et al.	06/17/1997	
	5	5,716,981		Hunter et al.	02/10/1998	
	6	5,733,876		O'Reilly et al.	03/31/1998	
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Examiner Signature	<i>S. Qazi</i>	Date Considered	2/23/06
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Sheet 2 of 15

## Complete if Known

Application No.	09/939,208
Filing Date	08/24/2001
First Named Inventor	Gregory E. Agoston et al.
Group Art Unit	1616
Examiner Name	Qazi, S.
Attorney Docket Number	05213-0852 (43170-263550)

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		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)			
	18	EPO	0166937	A2	Abbott Laboratories	08/01/1986	
	19	DE	27 57 157		Schering AG	12/19/1977	
SEA	20	WO	93/19746	A1	ORION-YH-TYMA OY	10/14/1993	
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	23	JP	58-131978		?	08/06/1983	No
	24	JP	63-119500		?	05/24/1988	No

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S. Qazi

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	25	Author: Title: <i>Incomplete</i> Publ: <i>Research Plus Catalog</i> Vol/Iss: Pages: 50-58 Date: 1993			
	26	Author: Title: Registry Number: 56933-77-8 Publ: <i>Chemical Abstracts</i> Vol/Iss: Pages: Date:			
	27	Author: Title: Registry Number: 56933-78-9 Publ: <i>Chemical Abstracts</i> Vol/Iss: Pages: Date:			
	28	Author: Title: Registry Number: 57380-15-1 Publ: <i>Chemical Abstracts</i> Vol/Iss: Pages: Date:			
<i>SB</i>	29	Author: Title: Registry Number: 71782-94-0 Publ: <i>Chemical Abstracts</i> Vol/Iss: Pages: Date:			
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	31	Author: Title: Registry Number: 101277-11-6 Publ: <i>Chemical Abstracts</i> Vol/Iss: Pages: Date:			

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	32	Author: Title: Registry Number: 101429-40-7 Publ: <u>Chemical Abstracts</u> Vol/Iss: Pages: Date:			
SA	33	Author: Title: Registry Number: 162853-20-5 Publ: <u>Chemical Abstracts</u> Vol/Iss: Pages: Date:			
	34	Author: Title: News Article: Hoffman-La Roche Signs \$70 Million Deal with Millenium on Genomics Technology Publ: <u>Genetic Engineering News</u> Vol/Iss: <u>Incomplete</u> Pages: Date: 4/15/1994			
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	36	Author: Title: News Article: Nasal Spray for Treating Bleeding Disorders Publ: <u>Genetic Engineering News</u> Vol/Iss: <u>Incomplete</u> Pages: Date: 4/15/1994			
SA	37	Author: ALGIRE, G.H. et al. Title: Vascular reactions of normal and malignant tumors in vivo. I. Vascular reactions of mice to wounds and to normal and neoplastic transplants Publ: <u>Journal of the National Cancer Institute</u> Vol/Iss: 6 Pages: 73-85 Date: 08/1945			
	38	Author: ALIEV et al. Title: 54929q Synthesis of cycloalkyl derivatives of dihydric phenols and their ethers Publ: <u>Chemical Abstracts</u> Vol/Iss: 72 Pages: 370 Date: 1970			

Examiner Signature	<i>S. Qazi</i>	Date Considered	2/23/06
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	39	Author: ANSTEAD et al. Title: The Estradiol Pharmacophore: Ligand Structure-Estrogen Receptor Binding Affinity Relationships and a Model for the Receptor Binding Site Publ: Steroids Vol/Iss: 62 Pages: 268-303 Date: 1997			
	40	Author: ATTALLA et al. Title: 2-Methoxyestradiol-Induced Phosphorylation of Bcl-2: Uncoupling from JNK/SAPK Activation (Abstract only) Publ: Biochemical and Biophysical Research Communications Vol/Iss: 247 (3) Pages: 616-619 Date: 6/29/1998			
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	42	Author: BINDRA et al. Title: Studies in Antifertility Agents. 8. Seco Steroids. 2. 5,6-Secoestradiol and Some Related Compounds Publ: Journal of Medicinal Chemistry Vol/Iss: 18 (9) Pages: 921-925 Date: 1975			
	43	Author: BOYCE et al. Title: Some Preliminary Synthetical Studies with 5,6,7,8-Tetra-hydro-8-methylindane -1,5-dione Publ: Unknown Vol/Iss: 7 Pages: 4547-4553 Date: 1960			
SB	44	Author: BREM, H. et al. Title: Interstitial chemotherapy with drug polymer implants for the treatment of recurrent gliomas Publ: Journal of Neurosurgery Vol/Iss: 74 Pages: 441-446 Date: 03/01/91			
	45	Author: BURROWS, N.P. Title: Thalidomide Modifies Disease Publ: British Medical Journal Vol/Iss: 307 (6909) Pages: 939-940 Date: 10/09/93			

Examiner Signature	SB	Date Considered	2/23/06
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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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	46	Author: CAMBIE et al. Title: Aromatic Steroids. Part I. Oxidation Products of 3-Methoxyestra-1,3,5(10)-triene- 17β-yl Acetate Publ: J. Chem. Soc. Vol/Iss: Pages: 2603-2608 Date: 1968			
	47	Author: COREY et al. Title: Applications of N,N-Dimethylhydrazones to Synthesis. Use in Efficient, Positionally and Stereochemically Selective C-C Bond Formation; Oxidative Hydrolysis to Carbonyl Compounds Publ: Tetrahedron Letters Vol/Iss: 1 Pages: 3-6 Date: 1976			
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	50	Author: DING et al. Title: Sex Hormone-Binding Globulin Mediates Prostate Androgen Receptor Action via a Novel Signaling Pathway (Abstract only) Publ: Endocrinology Vol/Iss: 139 (1) Pages: 213-218 Date: 1998			
	51	Author: DURANI et al. Title: Seco-Oestradiols and Some Non-Steroidal Oestrogens: Structural Correlates of Oestrogenic Action Publ: Journal of Steroid Biochemistry Vol/Iss: 11 Pages: 67-77 Date: 1979			
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Examiner Signature	Sabih Qazi	Date Considered	2/23/06
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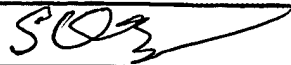
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	53	Author: FEVIG et al. Title: A Short, Stereoselective Route to 16 $\alpha$ -(Substituted-alkyl)estradiol Derivatives Publ: <u>Journal of Organic Chemistry</u> Vol/Iss: 52 Pages: 247-251 Date: 1987			
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	55	Author: FIESER et al. Title: N-Methylformanilide Publ: <u>Organic Synthesis Collective Volume 3</u> Vol/Iss: 3 Pages: 590-591 Date: 1955			
	56	Author: FOLKMAN, J. Title: Tumor Angiogenesis: Therapeutic Implications Publ: <u>New England Journal of Medicine</u> Vol/Iss: 285 (21) Pages: 1182-1186 Date: 11/18/71			
	57	Author: FOLKMAN, J. et al. Title: Induction of Angiogenesis During the Transition from Hyperplasia to Neoplasia Publ: <u>Nature</u> Vol/Iss: 339 Pages: 58-61 Date: 05/04/89			
	58	Author: FOLKMAN, J. et al. Title: Tumor Behavior in Isolated Perfused Organs in vitro Growth and Metastases of Biopsy Material in Rabbit Thyroid and Canine Intestinal Segment Publ: <u>Annals of Surgery</u> Vol/Iss: 164(3) Pages: 491-502 Date: 09/01/66			
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Examiner Signature	<i>S. Qazi</i>	Date Considered	2/3/06
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Sheet	8	of	15	Attorney Docket Number	
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OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS					
Examiner Initials	Cite No.	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published			T*
	60	Author: GASLINI et al. Title: Reaction of Eugenol with Synthesis Gas. Synthesis of 5,6,7,8-Tetrahydro-3-methoxy-2-naphthol Publ: Journal of Organic Chemistry Vol/Iss: 29 (5) Pages: 1177-1180 Date: 05/1984			
	61	Author: GIMBRONE, M.A. et al. Title: Tumor Growth and Neovascularization: An Experimental Model Using the Rabbit Cornea Publ: Journal of the National Cancer Institute Vol/Iss: 52(2) Pages: 413-427 Date: 02/74			
	62	Author: GIMBRONE, M.A. et al. Title: Tumor dormancy <i>in vivo</i> by Prevention of Neovascularization Publ: Journal of Experimental Medicine Vol/Iss: 136 Pages: 261-276 Date: 1972			
	63	Author: GONZALEZ et al. Title: Synthesis and Pharmacological Evaluation of 8 $\alpha$ Estradiol Derivatives Publ: Steroids Vol/Iss: 40 (2) Pages: 171-187 Date: 9/1982			
	64	Author: GROSS, J.L. et al. Title: Modulation of Solid Tumor Growth <i>in vivo</i> by bFGF (Abstract only) Publ: Proceedings of the American Association of Cancer Research Vol/Iss: 31 Pages: 79 Date: 03/90			
	65	Author: GUNZLER, V. Title: Thalidomide-A Therapy for the Immunological Consequences of HIV Infection? Publ: Medical Hypothesis Vol/Iss: 30 (2) Pages: 105-109 Date: 10/89			
	66	Author: GUPTA et al. Title: Antifertility Agents. XIV. Secosteroids. VII. Synthesis of 2 $\alpha$ and 2 $\beta$ , 6 $\beta$ -dimethyl- 3 $\beta$ -(p-hydroxyphenyl)-trans-bicyclo[4.3.0]nonan-7-ones and some related compounds (Abstract only) Publ: Indian Journal of Chemistry Vol/Iss: 13 (7) Pages: 759-760 Date: 1975			

Examiner Signature		Date Considered	2/23/05
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\*Unique citation designation number. \*See attached Kinds of U.S. Patent Documents. \*Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). \*For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. \*Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. \*Applicant is to place a check mark here if English language translation is attached.



Substitute for Form 1449/A/PTO				DEC 12 2003 <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)		Complete if Known	
Sheet 9 of 15				Application No.		09/939,208	
				Filing Date		08/24/2001	
				First Named Inventor		Gregory E. Agoston et al.	
				Group Art Unit		1616	
				Examiner Name		Qazi, S.	
				Attorney Docket Number		05213-0852 (43170-263550)	
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	67	Author: GUPTA et al. Title: Studies in Antifertility Agents. Part XVIII. 2α6β-Diethyl-3β-(p-hydroxyphenyl)-trans-bicyclo[4.3.0]nonan-7β-ol and 6β-methyl-3β-(p-hydroxyphenyl)-2α-propyl- trans-bicyclo[4.3.0]nonan-7β-ol (Abstract only) Publi: Indian Journal of Chemistry Vol/Iss: 19B (10) Pages: 886-890 Date: 1980					
	68	Author: GUTIERREZ-RODRIGUEZ et al. Title: Treatment of Refractory Rheumatoid Arthritis - The Thalidomide Experience Publi: The Journal of Rheumatology Vol/Iss: 16 (2) Pages: 158-163 Date: 02/89					
	69	Author: GUTIERREZ-RODRIGUEZ, O. Title: Thalidomide - A Promising New Treatment for Rheumatoid Arthritis Publi: Arthritis and Rheumatism Vol/Iss: 27 (10) Pages: 1118-1121 Date: 10/84					
	70	Author: HAN et al. Title: Dehydroepiandrosterone and Dihydrotestosterone Recognition by Human Estrogenic 17β-Hydroxysteroid Dehydrogenase Publi: Journal of Biological Chemistry Vol/Iss: 275 Iss 2 Pages: 1105-1111 Date: 1/14/2000					
	71	Author: HANDLEY et al. Title: Chronic bullous disease of childhood and ulcerative colitis Publi: British Journal of Dermatology Vol/Iss: 127 (40) Pages: 67-68 Date: 07/01/92					
	72	Author: HENEY et al. Title: Thalidomide treatment for chronic graft-versus-host disease Publi: British Journal of Haematology Vol/Iss: 78 (1) Pages: 23-27 Date: 05/91					
	73	Author: HORI, A. et al. Title: Suppression of Solid Tumor Growth by Immunoneutralizing Monoclonal Antibody Against Human Basic Fibroblasts Growth Factor Publi: Cancer Research Vol/Iss: 51 Pages: 6180-6184 Date: 11/15/91					

Examiner Signature		Date Considered	2/23/01
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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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				Application No.	09/939,208
				Filing Date	08/24/2001
				First Named Inventor	Gregory E. Agoston et al.
				Group Art Unit	1616
Sheet 10 of 15				Examiner Name	Qazi, S.
				Attorney Docket Number	05213-0852 (43170-263550)
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	74	Author: IMAMURA et al. Title: Method for Manufacture of Dihydric Phenols (Abstract only) Publ: USPATFULL 76:20259 US 3,950,437 Vol/Iss: Pages: Date: 04/13/1976			
	75	Author: INGBER, D. et al. Title: Synthetic analogues of fumagillin that inhibit angiogenesis and suppress tumor growth Publ: Nature Vol/Iss: 348 Pages: 555-557 Date: 12/06/90			
	76	Author: JHINGRAN et al. Title: Studies in Antifertility Agents - Part XLI: Secosteroids-x: Syntheses of Various Stereoisomers of (+)-2,6β-diethyl-7α-ethynyl-3-(p-hydroxyphenyl)-trans-bicyclo [4.3.0]nonan-7β-ol. Publ: Steroids Vol/Iss: 42 (6) Pages: 627-634 Date: 1983			
	77	Author: KARWAT Title: Separation and Recovery of Hydrogen Sulfide from Hydrocarbon Mixture Publ: Caplus DE 1103310 Vol/Iss: Pages: Date: 09/02/1959			
	78	Author: KATAOKA et al. Title: An Agent that Increases Tumor Suppressor Transgene Product Coupled with Systemic Transgene Delivery Inhibits Growth of Metastatic Lung Cancer in Vivo (Abstract only) Publ: Cancer Research Vol/Iss: 58 (21) Pages: 4761-4765 Date: 11/1998			
	79	Author: KIM, K.J. et al. Title: Inhibition of Vascular Endothelial Growth Factor-induced Angiogenesis Suppresses Tumor Growth In Vivo Publ: Nature Vol/Iss: 362 Pages: 841-844 Date: 04/29/93			
	80	Author: KNIGHTON, D. et al. Title: Avascular and Vascular Phases of Tumour Growth in the Chick Embryo Publ: British Journal of Cancer Vol/Iss: 35 Pages: 347-356 Date: 1977			

Examiner Signature	Sabiha Qazi	Date Considered	2/23/06
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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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**INFORMATION DISCLOSURE  
STATEMENT BY APPLICANT**

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Application No.	09/939,208
Filing Date	08/24/2001
First Named Inventor	Gregory E. Agoston et al.
Group Art Unit	1616
Examiner Name	Qazi, S.
Attorney Docket Number	05213-0852 (43170-263550)

Sheet

11

of

15

**OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS**

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	81	Author: KOLE et al. Title: Studies in Antifertility Agents. 11. Secosteroids. 5. Synthesis of 9,11-Secoestradiol Publ: <u>Journal of Medicinal Chemistry</u> Vol/Iss: 18 (7) Pages: 765-766 Date: 1975	
	82	Author: LEWIS, Richard J. Title: Publ: <u>Hawley's Condensed Chemical Dictionary</u> Vol/Iss: Pages: 577 Date: 01/1993	
	83	Author: LEWIS, Richard J. Title: Publ: <u>Hawley's Condensed Chemical Dictionary</u> Vol/Iss: Pages: 128-129 Date: 01/1993	
SOZ	84	Author: LIEN, W. et al. Title: The blood supply of experimental liver metastases. II. A Microcirculatory study of the normal and tumor vessels of the liver with the use of perfused silicone rubber Publ: <u>Surgery</u> Vol/Iss: 68 (2) Pages: 334-340 Date: 08/1970	
	85	Author: LIU et al. Title: Total Synthesis of (+)- $\Delta^{9(12)}$ Capnellene Publ: <u>Tetrahedron Letters</u> Vol/Iss: 26 (40) Pages: 4847-4850 Date: 1985	
	86	Author: MARO et al. Title: Mechanism of Polar Body Formation in the Mouse Oocyte: An Interaction Between the Chromosomes, the Cytoskeleton and the Plasma Membrane Publ: <u>Journal of Embryology and Experimental Morphology</u> Vol/Iss: 92 Pages: 11-32 Date: 1986	
	87	Author: MEZA et al. Title: Managing the Gastrointestinal Complications of AIDS Publ: <u>Drug Therapy</u> Vol/Iss: 23 (11) Pages: 74-83 Date: 11/1993	

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2/23/06

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	88	Author: NAAFS et al. Title: Thalidomide Therapy An Open Trial Publ: International Journal of Dermatology Vol/Iss: 24 (2) Pages: 131-134 Date: 03/1985			
	89	Author: NAMBARA et al. Title: Synthesis of 16 $\alpha$ -Oxygenated Catechol Estrogen Methyl Ethers, New and Potential Metabolites Publ: Chemical & Pharmaceutical Bulletin Vol/Iss: 23 (7) Pages: 1613-1616 Date: 07/1975			
	90	Author: NEWKOME et al. Title: Synthesis of Simple Hydrazones of Carbonyl Compounds by an Exchange Reaction Publ: Journal of Organic Chemistry Vol/Iss: 31 Pages: 677-681 Date: 03/1966			
	91	Author: NGUYEN, M. et al. Title: Elevated Levels of the Angiogenic Peptide Basic Fibroblast Growth Factor in Urine of Bladder Cancer Patients Publ: Journal of the National Cancer Institute Vol/Iss: 85 (3) Pages: 241-242 Date: 02/03/93			
	92	Author: PAKALA et al. Title: Modulation of Endothelial Cell Proliferation by Retinoid x Receptor Agonists Publ: European Journal of Pharmacology Vol/Iss: 385 (2/3) Pages: 255-261 Date: 09/1999			
	93	Author: PENG et al. Title: Synthesis and Optical Properties of Novel Unsymmetrical Conjugated Dendrimers Publ: Journal of the American Chemical Society Vol/Iss: 122 Pages: 6619-6623 Date: 2000			
	94	Author: POWELL et al. Title: Investigation and Treatment of Orogenital Ulceration; studies on a Possible Mode of Action of Thalidomide Publ: British Journal of Dermatology Vol/Iss: 113 Supp. 28 Pages: 141-144 Date: 07/1985			

Examiner Signature	<i>S. Qazi</i>	Date Considered	2/23/06
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	95	Author: ROMANELLI et al. Title: Ethyl-p-Dimethylaminophenylacetate Publ: Organic Synthesis Vol/Iss: 5 Pages: 552 Date:			
	96	Author: SEEGERs et al. Title: The Mammalian Metabolite, 2-methoxyestradiol, Affects P53 Levels and Apoptosis Induction in Transformed Cells but Not in Normal Cells (Abstract only) Publ: Journal of Steroid Biochemistry and Molecular Biology Vol/Iss: 62 (4) Pages: 253-267 Date: 07/1997			
	97	Author: SHAH et al. Title: (+/-)-(N-alkylamino)benzazepine Analogs: Novel Dopamine D1 Receptor Antagonists Publ: Journal of Medicinal Chemistry Vol/Iss: 38 (21) Pages: 4284-4293 Date: 10/13/95			
	98	Author: SIDKY et al. Title: Inhibition of Angiogenesis by Interferons: Effects on Tumor- and Lymphocyte-induced Vascular Responses Publ: Cancer Research Vol/Iss: 47 Pages: 5155-5161 Date: 10/01/87			
	99	Author: SRIVASTAVA, A. et al. Title: The Prognostic Significance of Tumor Vascularity in Intermediate-Thickness (0.76-4.0 mm Thick) Skin Melanoma Publ: American Journal of Pathology Vol/Iss: 133 (2) Pages: 419-424 Date: 11/1988			
	100	Author: STARKOV et al. Title: Mono- and Dialkylation of Guaiacol by Olefins on KU-2 Cation Exchanger (Abstract only) Publ: Zhurnal Prikladnoi Khimii Vol/Iss: 41 (3) Pages: 688-690 Date: 1968			
	101	Author: TAYLOR, S. et al. Title: Protamine is an Inhibitor of Angiogenesis Publ: Nature Vol/Iss: 297 Pages: 307-312 Date: 05/27/82			

Examiner Signature	Sabiha Qazi	Date Considered	2/23/06
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	102	Author: TREMBLAY et al. Title: A Convenient Synthetic Method for Alpha-Alkylation of Steroidal 17-Ketone: Preparation of 16 $\beta$ -(THPO-Heptyl)-Estradiol Publ: <u>Synthetic Communications</u> Vol/Iss: 25 (16) Pages: 2483-2495 Date: 1995					
	103	Author: TREMBLAY et al. Title: Synthesis of 16-(Bromoalkyl)-Estradiols Having Inhibitory Effect on Human Placental Estradiol 17 $\beta$ -Hydroxysteroid Dehydrogenase (17 $\beta$ HSD Type 1) Publ: <u>Bioorganic &amp; Medicinal Chemistry</u> Vol/Iss: 3 (5) Pages: 505-523 Date: 1995					
	104	Author: VICENTE et al. Title: In Vitro Activity of Thalidomide Against Mycobacterium avium Complex Publ: <u>Archives of Internal Medicine</u> Vol/Iss: 153 (4) Pages: 534 Date: 02/22/93					
	105	Author: WEIDNER, N. et al. Title: Tumor angiogenesis: A New Significant and Independent Prognostic Indicator in Early-Stage Breast Carcinoma Publ: <u>Journal of the National Cancer Institute</u> Vol/Iss: 84 Pages: 1875-1887 Date: 12/16/92					
	106	Author: WEIDNER, N. et al. Title: Tumor Angiogenesis Correlates with Metastasis in Invasive Prostate Carcinoma Publ: <u>American Journal of Pathology</u> Vol/Iss: 143 (2) Pages: 401-409 Date: 08/1993					
	107	Author: WEIDNER, N. et al. Title: Tumor Angiogenesis and Metastasis-Correlation in Invasive Breast Carcinoma Publ: <u>New England Journal of Medicine</u> Vol/Iss: 324 (1) Pages: 1-8 Date: 01/03/91					
	108	Author: WELSCH et al. Title: Staphylostatic Activity of Some New Diphenols, Naphthols, and Chalcones (Abstract only) Publ: <u>Experientia</u> Vol/Iss: 11 Pages: 350-351 Date: 1955					

Examiner Signature		Date Considered	2/23/06
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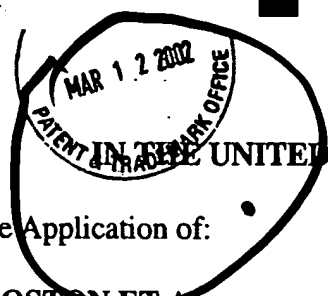
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S B 9	109	Author: WHITE et al. Title: Treatment of Pulmonary Hemangiomatosis with Recombinant Interferon Alfa-2a Publ: The New England Journal of Medicine Vol/Iss: 32 (18) Pages: 1197-1200 Date: 05/04/89				
	110	Author: ZOUBINE et al. Title: 2-Methoxyestradiol-Induced Growth Suppression and Lethality in Estrogen-Responsive MCF-7 Cells May Be Mediated by Down Regulation of p34cdc2 and Cyclin B1 Expression (Abstract only) Publ: International Journal of Oncology Vol/Iss: 15 (4) Pages: 639-646 Date: 10/1999				

Examiner Signature	S B 9	Date Considered	2/23/06
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UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of: )  
AGOSTON ET AL. ) Group Art No: 1623  
Serial No: 09/939,208 )  
Filed: August 24, 2001 )  
For: ANTIANGIOGENIC AGENTS )

INFORMATION DISCLOSURE STATEMENT

Assistant Commissioner for Patents  
Washington, DC 20231

*No page 19 in IDS*

Sir:

The citation of information on the attached Form PTO/SB/08A, "List of Art Cited by Applicant" is made pursuant to 37 C.F.R. §§ 1.97 and 1.98. A copy of all listed items is enclosed. The citation of this information does not constitute an admission that any of the materials are available as a reference or of priority, or a waiver of any right applicant may have under applicable statutes, Rules of Practice in patent cases, or otherwise.

Related applications are listed on the attached Appendix A for your reference. Applicants respectfully request that the Examiner review the applications listed in Appendix A, initial Appendix A in the appropriate spaces, and return a copy of the initialized Appendix to Applicants.

KILPATRICK STOCKTON LLP  
Suite 2800  
1100 Peachtree Road  
Atlanta, Georgia 30309  
(404) 815-6500  
Our Docket: 05213-0852 (43170-263550)

Respectfully submitted,  
*Suzanne Seavello Shope*  
By: Suzanne Seavello Shope  
Reg. No. 37,933

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: Assistant Commissioner for Patents, Washington, DC, 20231, on this 5th day of March 2002.

*Suzanne Seavello Shope*  
Suzanne Seavello Shope - Reg. No. 37,933